

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DRUGB, DRUGLAUNCH, DRUGMONOG2, ...' ENTERED AT 10:35:16 ON 07 APR 2003

SEA IDURONIDASE OR ALDURAZYME

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3   FILE ADISINSIGHT
9   FILE ADISNEWS
5   FILE AGRICOLA
3   FILE ANABSTR
1   FILE AQUASCI
5   FILE BIOBUSINESS
17  FILE BIOCOMMERCE
408 FILE BIOSIS
32  FILE BIOTECHABS
32  FILE BIOTECHDS
150 FILE BIOTECHNO
22  FILE CABA
38  FILE CANCERLIT
276 FILE CAPLUS
5   FILE CEABA-VTB
44  FILE CIN
11  FILE CONFSCI
13  FILE DDFB
17  FILE DDFU
67  FILE DGENE
13  FILE DRUGB
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6   FILE DRUGUPDATES
4   FILE EMBAL
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77  FILE ESBIODBASE
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49  FILE PHIN
168 FILE PROMT
303 FILE SCISEARCH
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132 FILE USPATFULL
2   FILE USPAT2
1   FILE VETU
26  FILE WPIDS
26  FILE WPINDEX

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L2

QUE IDURONIDASE OR ALDURAZYME

FILE 'BIOSIS, EMBASE, MEDLINE, SCISEARCH, CAPLUS, PROMT, BIOTECHNO, USPATFULL, PASCAL' ENTERED AT 10:37:03 ON 07 APR 2003

L3

6 S L1 AND MUTEIN

L4

6 S L2 AND MUTEIN

L5

6 DUP REM L4 (0 DUPLICATES REMOVED)

=> d 15 ibib ab 1-6

L5 ANSWER 1 OF 6 USPATFULL

ACCESSION NUMBER: 2003:51224 USPATFULL  
TITLE: Peptide extended glycosylated polypeptides  
INVENTOR(S): Okkels, Jens Sigurd, Vedbaek, DENMARK  
Jensen, Anne Dam, Copenhagen, DENMARK  
van den Hazel, Bart, Copenhagen, DENMARK

	NUMBER	KIND	DATE
PATENT INFORMATION:	US 2003036181	A1	20030220
APPLICATION INFO.:	US 2001-896896	A1	20010629 (9)

	NUMBER	DATE
PRIORITY INFORMATION:	DK 2000-1027	20000630
	DK 2000-1092	20000714
	WO 2000-DK743	20001229
	WO 2001-DK90	20010209
	US 2000-217497P	20000711 (60)
	US 2000-225558P	20000816 (60)

DOCUMENT TYPE: Utility  
FILE SEGMENT: APPLICATION  
LEGAL REPRESENTATIVE: MAXYGEN, INC., 515 GALVESTON DRIVE, RED WOOD CITY, CA, 94063

NUMBER OF CLAIMS: 57  
EXEMPLARY CLAIM: 1  
NUMBER OF DRAWINGS: 2 Drawing Page(s)  
LINE COUNT: 4732

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Glycosylated polypeptides comprising the primary structure NH.sub.2--X--Pp--COOH, wherein X is a peptide addition comprising or contributing to a glycosylation site, and Pp is a polypeptide of interest or comprising the primary structure NH.sub.2-P.sub.x--X--P.sub.y-COOH, wherein P.sub.x is an N-terminal part of a polypeptide Pp of interest, P.sub.y is a C-terminal part of said polypeptide Pp, and X is a peptide addition comprising or contributing to a glycosylation site are provided. The glycosylated polypeptides possess improved properties as compared to the polypeptide of interest.

L5 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:368665 CAPLUS  
DOCUMENT NUMBER: 136:385047  
TITLE: Methods for large scale production and purification of human .alpha.-L-iduronidase for treatment of mucopolysaccharidosis I  
INVENTOR(S): Qin, Minmin; Chan, Wai-Pan; Chen, Lin; Fitzpatrick, Paul A.; Henstrand, John M.; Wendt, Dan J.; Zecherle, Gary N.; Starr, Christopher M.; Kakkis, Emil D.  
PATENT ASSIGNEE(S): Biomarin Pharmaceutical, Inc., USA  
SOURCE: PCT Int. Appl., 71 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 3  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002038775	A2	20020516	WO 2001-US47843	20011109
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,				

GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,  
 LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,  
 RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,  
 UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,  
 BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

AU 2002027369 A5 20020521 AU 2002-27369 20011109

US 2002146802 A1 20021010 US 2001-993038 20011113

PRIORITY APPLN. INFO.:

US 2000-711202 A 20001109

US 1999-439923 A2 19991112

WO 2001-US47843 W 20011109

AB The present invention provides a recombinant human .alpha.-L-  
**iduronidase** and biol. active fragments and **muteins**  
 thereof with a purity greater than 99%. The present invention further  
 provides large-scale methods to produce and purify com. grade recombinant  
 human .alpha.-L-**iduronidase** enzyme thereof. The method involves  
 prepn. of a seed culture contg. Chinese hamster ovary cells 2.131  
 transfected with a vector encoding cDNA for .alpha.-L-**iduronidase**  
 . These cells is washed and resuspended in a protein-free culture medium  
 supplemented with 7.6 mg/L thymidine, 13.6 mg/L hypoxanthine, 375 .mu.g/mL  
 G418 and 5% fetal bovine serum. The cell suspension is incubated at  
 37.degree.C for 2-3 days with 5% CO2 in three 225 cm flasks. The said  
 cell suspension is split by sequentially adding the cells to one 1L  
 spinner flask, two 3L flasks and 4 8L flasks. The cell suspension is  
 stirred at 50 rpm, followed by increasing the inoculum vol. by incubating  
 and subculturing cells to a final cell d. of about 2-2.5 x 105. A mixt.  
 contg. macroporous microcarriers is prepd. in growth medium with fetal  
 bovine serum and transferring said mixt. to a bioreactor. Cells from the  
 bioreactor may be harvested at a d. of about 106. Methods for purifn. of  
 .alpha.-L-**iduronidase** to greater than 99% purity include  
 adjusting the pH to an acidic range, followed by filtering the mixt.  
 through a 0.2-0.54 .mu. filter. The filtrate is further passed through a  
 blue sepharose FF column to capture the protein which purifies .alpha.-L-  
**iduronidase** 7-10-fold. Contaminating CHO proteins are removed by  
 passing the fluid through a copper chelating sepharose column. The mixt.  
 is then passed through a Ph sepharose column to reduce residual leached  
 Cibacron blue dye and copper ions carried over from the previous columns.  
 Purified .alpha.-L-**iduronidase** is concd. and diafiltered. The  
 purifn. steps include 10% glycerol in all buffers to improve the  
 .alpha.-L-**iduronidase** yield. The specific activity of  
 .alpha.-L-**iduronidase** may be greater than 240,000 units/mg  
 protein.

L5 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: 2002:368334 CAPLUS

DOCUMENT NUMBER: 136:363851

TITLE: Methods to mass produce recombinant human .alpha.-L-  
**iduronidase** for treating diseases caused by  
 .alpha.-L-**iduronidase** deficiencies

INVENTOR(S): Kakkis, Emil D.

PATENT ASSIGNEE(S): Biomarin Pharmaceutical, Inc., USA

SOURCE: PCT Int. Appl., 73 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002038171	A2	20020516	WO 2001-US47835	20011109

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,  
 CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,

HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,  
 LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,  
 RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,  
 VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,  
 BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

AU 2002028991 A5 20020521 AU 2002-28991 20011109  
 US 2002164758 A1 20021107 US 2001-993241 20011113

PRIORITY APPLN. INFO.: US 2000-711205 A 20001109  
 US 1999-439923 A2 19991112  
 WO 2001-US47835 W 20011109

AB The present invention provides a method to mass produce human recombinant .alpha.-L-**iduronidase** in large scale amts. with appropriate purity to enable large scale prodn. for long term patient use of the enzyme therapy. The present invention provides a formulation comprising a pharmaceutical compn. comprising a human recombinant .alpha.-L-**iduronidase** or biol. active or **muteins** thereof with a purity of greater than 99%, or in combination with a pharmaceutically acceptable carrier. The present invention further provides methods to treat certain genetic disorders including .alpha.-L-**iduronidase** deficiency and mucopolysaccharidosis I (MPS 1) by administering said formulation.

L5 ANSWER 4 OF 6 USPATFULL

ACCESSION NUMBER: 2002:294715 USPATFULL  
 TITLE: Methods for treating diseases caused by deficiencies of recombinant alpha-L-**iduronidase**  
 INVENTOR(S): Kakkis, Emil D., Novato, CA, UNITED STATES

	NUMBER	KIND	DATE
PATENT INFORMATION:	US 2002164758	A1	20021107
APPLICATION INFO.:	US 2001-993241	A1	20011113 (9)
RELATED APPLN. INFO.:	Continuation of Ser. No. US 2000-711205, filed on 9 Nov 2000, PENDING Continuation-in-part of Ser. No. US 1999-439923, filed on 12 Nov 1999, PENDING		
DOCUMENT TYPE:	Utility		
FILE SEGMENT:	APPLICATION		
LEGAL REPRESENTATIVE:	HOWREY SIMON ARNOLD & WHITE, LLP, BOX 34, 301 RAVENSWOOD AVE., MENLO PARK, CA, 94025		
NUMBER OF CLAIMS:	28		
EXEMPLARY CLAIM:	1		
NUMBER OF DRAWINGS:	20 Drawing Page(s)		
LINE COUNT:	2003		

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides a formulation comprising a pharmaceutical composition comprising a human recombinant .alpha.-L-**iduronidase** or biologically active or **muteins** thereof with a purity of greater than 99%, or in combination with a pharmaceutically acceptable carrier. The present invention further provides methods to treat certain genetic disorders including .alpha.-L-**iduronidase** deficiency and mucopolysaccharidosis I (MPS 1) by administering said formulation.

L5 ANSWER 5 OF 6 USPATFULL

ACCESSION NUMBER: 2002:265926 USPATFULL  
 TITLE: Methods for producing and purifying recombinant alpha-L-**iduronidase**  
 INVENTOR(S): Qin, Minmin, Pleasanton, CA, UNITED STATES  
 Chan, Wai-Pan, Castro Valley, CA, UNITED STATES  
 Chen, Lin, San Francisco, CA, UNITED STATES  
 Fitzpatrick, Paul A., Albany, CA, UNITED STATES  
 Henstrand, John M., Davis, CA, UNITED STATES  
 Wendt, Dan J., Walnut Creek, CA, UNITED STATES

Zecherle, Gary N., Novato, CA, UNITED STATES  
Starr, Christopher M., Sonoma, CA, UNITED STATES  
Kakkis, Emil D., Novato, CA, UNITED STATES

	NUMBER	KIND	DATE
PATENT INFORMATION:	US 2002146802	A1	20021010
APPLICATION INFO.:	US 2001-993038	A1	20011113 (9)
RELATED APPLN. INFO.:	Continuation of Ser. No. US 2000-711202, filed on 9 Nov 2000, PENDING Continuation-in-part of Ser. No. US 1999-439923, filed on 12 Nov 1999, PENDING		
DOCUMENT TYPE:	Utility		
FILE SEGMENT:	APPLICATION		
LEGAL REPRESENTATIVE:	HOWREY SIMON ARNOLD & WHITE, LLP, BOX 34, 301 RAVENSWOOD AVE., MENLO PARK, CA, 94025		
NUMBER OF CLAIMS:	7		
EXEMPLARY CLAIM:	1		
NUMBER OF DRAWINGS:	19 Drawing Page(s)		
LINE COUNT:	1964		
CAS INDEXING IS AVAILABLE FOR THIS PATENT.			

AB The present invention provides a recombinant human .alpha.-L-iduronidase and biologically active fragments and muteins thereof with a purity greater than 99%. The present invention further provides large-scale methods to produce and purify commercial grade recombinant human .alpha.-L-iduronidase enzyme thereof.

L5 ANSWER 6 OF 6 USPATFULL

ACCESSION NUMBER: 2002:235036 USPATFULL  
TITLE: Lysosomal enzymes and lysosomal enzyme activators  
INVENTOR(S): Okkels, Jens Sigurd, Vedbaek, DENMARK  
Jensen, Anne Dam, Copenhagen, DENMARK  
Halkier, Torben, Solroed Strand, DENMARK  
Jensen, Rikke Bolding, Skibby, DENMARK  
Schambye, Hans Thalsgard, Frederiksberg, DENMARK

	NUMBER	KIND	DATE
PATENT INFORMATION:	US 2002127219	A1	20020912
APPLICATION INFO.:	US 2000-753126	A1	20001229 (9)

	NUMBER	DATE
PRIORITY INFORMATION:	DK 1999-1891	19991230
	DK 2000-865	20000602
	DK 2000-866	20000602
	DK 2000-1027	20000630
	US 2000-174652P	20000106 (60)
	US 2000-210984P	20000612 (60)
	US 2000-211124P	20000612 (60)
	US 2000-217497P	20000711 (60)

DOCUMENT TYPE: Utility  
FILE SEGMENT: APPLICATION  
LEGAL REPRESENTATIVE: LAW OFFICES OF JONATHAN ALAN QUINE, P O BOX 458, ALAMEDA, CA, 94501  
NUMBER OF CLAIMS: 58  
EXEMPLARY CLAIM: 1  
NUMBER OF DRAWINGS: 9 Drawing Page(s)  
LINE COUNT: 4771  
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A polypeptide selected from the group of lysosomal enzymes and lysosomal enzyme activators, comprising at least one introduced glycosylation site as compared to a corresponding parent enzyme or activator. By introducing additional glycosylation sites the resulting glycosylated

lysosomal enzyme or activator obtains improved in vivo activity and thereby provides for improved treatment of lysosomal storage diseases.

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS  
RN 9073-56-7 REGISTRY  
CN Iduronidase, .alpha.-L- (9CI) (CA INDEX NAME)  
OTHER NAMES:  
CN .alpha.-Iduronidase  
CN .alpha.-L-Iduronidase  
CN Aldurazyme  
CN E.C. 3.2.1.76  
MF Unspecified  
CI MAN  
LC STN Files: ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, CA, CAPLUS,  
CBNB, CIN, IPA, MRCK\*, PROMT, TOXCENTER, USPATFULL  
(\*File contains numerically searchable property data)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

260 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

260 REFERENCES IN FILE CAPLUS (1962 TO DATE)